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| Substitute Form PTO-1449<br>(Modified)   | U.S. Department of Commerce<br>Patent and Trademark Office | Attorney's Docket No.<br>14539-006002 | Application No.<br><u>10/625,105</u> |
| <b>Information Disclosure Statement<br/>by Applicant</b><br>(Use several sheets if necessary)<br>(37 CFR §1.98(b)) <u>07/22/2003</u> |  | Applicant<br>Takashi Tsuji et al.     |                                      |
|  |  | Filing Date<br>July 22, 2003          | Group Art Unit<br><u>1644</u>        |

| U.S. Patent Documents |           |                 |                  |                     |       |          |                            |
|-----------------------|-----------|-----------------|------------------|---------------------|-------|----------|----------------------------|
| Examiner Initial      | Desig. ID | Document Number | Publication Date | Patentee            | Class | Subclass | Filing Date If Appropriate |
| JO                    | AA        | 5,506,126       | 4/9/1996         | Seed et al.         |       |          |                            |
|                       | AB        | 5,521,288       | 5/28/1996        | Linsley et al.      |       |          |                            |
|                       | AC        | 6,075,181       | 6/13/2000        | Kucherlapati et al. |       |          |                            |
|                       | AD        | 2002/0156242    | 10/24/2002       | Tamatani et al.     |       |          |                            |

| Foreign Patent Documents or Published Foreign Patent Applications |           |                 |                  |                          |       |          |                            |
|---|-----------|-----------------|------------------|--------------------------|-------|----------|----------------------------|
| Examiner Initial  | Desig. ID | Document Number | Publication Date | Country or Patent Office | Class | Subclass | Translation<br>Yes      No |
|   | AE        | WO 98/11909     | 03/26/1998       | WIPO                     |       |          |                            |
|   | AF        | WO 98/38216     | 09/03/1998       | WIPO                     |       |          |                            |
|   | AG        | WO 99/15553     | 04/01/1999       | WIPO                     |       |          |                            |
|   | AH        | WO 00/19988     | 04/13/2000       | WIPO                     |       |          |                            |
|   | AI        | WO 00/46240     | 08/10/2000       | WIPO                     |       |          |                            |
|   | AJ        | WO 00/67788     | 11/16/2000       | WIPO                     |       |          |                            |
|   | AK        | WO 01/08700     | 02/08/2001       | WIPO                     |       |          |                            |
|   | AL        | WO 01/12658     | 02/22/2001       | WIPO                     |       |          |                            |
|   | AM        | WO 01/15732     | 03/08/2001       | WIPO                     |       |          |                            |
|   | AN        | WO 01/18022     | 03/15/2001       | WIPO                     |       |          |                            |
|   | AO        | WO 01/21796     | 03/29/2001       | WIPO                     |       |          |                            |
|   | AP        | WO 01/32675     | 05/10/2001       | WIPO                     |       |          |                            |
|   | AQ        | WO 01/64704     | 09/07/2001       | WIPO                     |       |          |                            |
|   | AR        | WO 01/87981     | 11/22/2001       | WIPO                     |       |          |                            |
|   | AS        | WO 02/44364     | 06/06/2002       | WIPO                     |       |          |                            |
|   | AT        | WO 02/70010     | 09/12/2002       | WIPO                     |       |          |                            |
|   | AU        | WO 02/76504     | 10/03/2002       | WIPO                     |       |          |                            |
|   | AV        | AU 13320/99     | 04/12/1999       | AU                       |       |          |                            |
|   | AW        | DE 19821060     | 04/15/1999       | DE                       |       |          |                            |
| JO  | AX        | EP 0984023 A1   | 03/08/2000       | EP                       |       |          |                            |

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|---|--------------------------------------|
| Examiner Signature<br><i>Ivan Ongspeslin</i>  | Date Considered<br><u>11/07/2005</u> |
| EXAMINER: Initials citation considered. Drawline through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |                                      |

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|---|-----------|-----------------|------------------|--------------------------|-------|----------|-------------|-----|----|
| Examiner Initial  | Desig. ID | Document Number | Publication Date | Country or Patent Office | Class | Subclass | Translation | Yes | No |
| <u>JG</u>   | AY        | EP 1 125 585 A1 | 08/22/2001       | EP                       |       |          |             |     |    |

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|   | AAA       | Bajorath, "A molecular model of inducible costimulator protein and three-dimensional analysis of its relation to the CD28 family of T cell-specific costimulatory receptors," J. MOL. MODEL 5:169-176 (1999)                                   |
|   | ABB       | Beier et al., "Induction, binding specificity and function of human ICOS," EUR. J. IMMUNOL., 30(12):3707-3717 (2000)   |
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|   | AFF       | Cameron "Recent advances in transgenic technology" MOLECULAR BIOTECHNOLOGY 7:253-65 (1997)   |
|   | AGG       | Chambers, "The expanding world of co-stimulation: the two-signal model revisited," TRENDS IN IMMUNOLOGY, 22(4):217-223 (2001)  |
|   | AHH       | Cocks et al., "A novel receptor involved in T-cell activation," NATURE, 376:260-263 (1995)   |
|   | AII       | Coyle et al., "The CD28-Related Molecule ICOS Is Required for Effective T Cell-Dependent Immune Responses," IMMUNITY 13(1):95-105 (2000)   |
|   | AJJ       | Dong et al., "Cutting Edge: Critical Role of Inducible Costimulator in Germinal Center Reactions," J. IMMUNOL., 166(6):3659-3662 (2001)  |
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|   | AMM       | Guo et al., "Stimulatory Effects of B7-Related Protein-1 on Cellular and Humoral Immune Responses in Mice," J. IMMUNOL., 166(9):5578-5584 (2001)   |
|   | ANN       | Hanzawa et al., "Characteristics of a TTH1 antibody which blocks an unknown adhesion phenomenon," PROCEEDINGS OF THE JAPANESE SOCIETY FOR IMMUNOLOGY, Vol. 24, Abstract No. W17-13 (1994) [ORIGINAL JAPANESE AND ENGLISH LANGUAGE TRANSLATION] |
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|   | ATT       | Kopf et al., "Inducible Costimulator Protein (ICOS) Controls T Helper Cell Subset Polarization after Virus and Parasite Infection," J. EXP. MED., 192(1):53-61 (2000)   |  |
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|   | ACCC      | Mullins et al. "Expression of the DBA/2J Ren-2 gene in the adrenal gland of transgenic mice" EMBO J., 8:4065-72 (1989)  |  |
|   | ADDD      | Mullins et al. "Fulminant hypertension in transgenic rats harbouring the mouse Ren-2 gene" NATURE, 344:541-44 (1990)  |  |
|   | AEEE      | Mullins et al. "Transgenesis in nonmurine species" Hypertension 22:630-33 (1993)  |  |
|   | AFFF      | Niemann "Transgenic farm animals get off the ground" TRANSGENIC RESEARCH, 7:73-75 (1998)  |  |
|   | AGGG      | Nojima et al., "The 4F9 antigen is a member of the tetra spans transmembrane protein family and functions as an accessory molecule in T cell activation and adhesion," CELLULAR IMMUNOLOGY, 152:249-260 (1993)                    |  |
|   | AHHH      | Overbeek "Factors affecting transgenic animal production," Transgenic Animal Technology, A Laboratory Handbook 96-98 (1994)   |  |
| <u>JO</u>   | AIII      | Özkaraynak et al., "Importance of ICOS-B7RP-1 costimulation in acute and chronic allograft rejection," NATURE IMMUNOLOGY 2(7):591-596 (2001)  |  |

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| Examiner Signature   | Date Considered   |
| <i>Ilia Onyschenko</i>   | <u>11/07/2005</u> |
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| Other Documents (include Author, Title, Date, and Place of Publication) |           |  |  |
|---|-----------|--|--|
| Examiner Initial  | Desig. ID | Document   |  |
| JO  | AJJJ      | Poster, Kyoto International Conference Hall, Takaragaike Sakyo-ku, Kyoto, JAPAN (November 30, 1994) [ORIGINAL JAPANESE AND ENGLISH LANGUAGE TRANSLATION]   |  |
|   | AKKK      | Redoglia et al., "Characterization of H4: a mouse T lymphocyte activation molecule functionally associated with the CD3/T cell receptor," EUR. J. IMMUNOL., 26(11):2781-2789 (1996)  |  |
|   | ALLL      | Riley et al., "ICOS Costimulation Requires IL-2 and Can Be Presented by CTLA-4 Engagement," J. IMMUNOL., 166(8):4943-4948 (2001)   |  |
|   | AMMM      | Robert et al., "Antibody Cross-Linking of the Thymocyte-Specific Cell Surface Molecule CTX Causes Abnormal Mitosis and Multinucleation of Tumor Cells," EXPERIMENTAL CELL RESEARCH, 235:227-237 (1997)   |  |
|   | ANNN      | Sato et al. (2000) "Up-regulation of inducible co-stimulator (ICOS) expression and its regulation of cytokine production in inflammatory bowel disease," GASTROENTEROLOGY, 118(4):A662   |  |
|   | A000      | Sharpe, "Analysis of lymphocyte costimulation <i>in vivo</i> using transgenic and 'knockout' mice," CURRENT OPINION IN IMMUNOLOGY, 7:389-395 (1995)  |  |
|   | APPP      | Sigmund "Are studies in genetically altered mice out of control?" ARTERIOSCLER. THROMB. VASC. BIOL., 20:1425-29 (2000)   |  |
|   | AQQQ      | Swallow et al., "B7h, a Novel Costimulatory Homolog of B7.1 and B7.2, Is Induced by TNF $\alpha$ ," IMMUNITY, 11(4):423-432 (1999)   |  |
|   | ARRR      | Tafuri et al., "ICOS is essential for effective T-helper-cell responses," NATURE 409(6816):105-109 (2001)  |  |
|   | ASSS      | Tai et al., "A role for CD9 molecules in T cell activation," J. EXP. MED., 184:753-758 (1996)  |  |
|   | ATTT      | Tamatani et al., "AILIM/ICOS: a novel lymphocyte adhesion molecule," INTERNATIONAL IMMUNOLOGY, 12(1):51-55 (2000)  |  |
|   | AUUU      | Tamatani et al., "Characteristics of an antibody which induces an ICAM-1-LFA-1-independent adhesion pathway," PROCEEDINGS OF THE JAPANESE SOCIETY FOR IMMUNOLOGY, Vol. 23, Abstract No. H-160 (1993) [ORIGINAL JAPANESE AND ENGLISH LANGUAGE TRANSLATION]                        |  |
|   | AVVV      | Tezuka et al., "Genetic cloning of a lymphocyte surface signal transduction molecule which induces an unknown adhesion phenomenon," PROCEEDINGS OF THE JAPANESE SOCIETY FOR IMMUNOLOGY, Vol. 24, Abstract No. W17-14 (1994) [ORIGINAL JAPANESE AND ENGLISH LANGUAGE TRANSLATION] |  |
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|   | AXXX      | Wall "Transgenic livestock: progress and prospects for the future" THERIOGENOLOGY 45:S7-68 (1996)  |  |
|   | AYYY      | Wang et al., "Costimulation of T cells by B7-H2, a B7-like molecule that binds ICOS," BLOOD, 96(8):2808-2813 (2000)  |  |
|   | AZZZ      | Yoshinaga et al., "Characterization of a new human B7-related protein: B7RP-1 is the ligand to the co-stimulatory protein ICOS," INTERNATIONAL IMMUNOLOGY, 12(10):1439-1447 (2000)   |  |
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Sheet 1 of 3

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| U.S. Patent Documents |           |                 |                  |                 |       |          |                            |
|-----------------------|-----------|-----------------|------------------|-----------------|-------|----------|----------------------------|
| Examiner Initial      | Desig. ID | Document Number | Publication Date | Patentee        | Class | Subclass | Filing Date If Appropriate |
| J0                    | AA        | 5,484,892       | 01/16/1996       | Tedder et al.   |       |          |                            |
|                       | AB        | 5,747,461       | 05/05/1998       | Markov          |       |          |                            |
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|                       | AF        | 20020115831     | 08/22/2002       | Tamatani et al. |       |          |                            |
|                       | AG        | 20020164697     | 11/07/2002       | Coyle et al.    |       |          |                            |
|                       | AH        | 20020177191     | 11/28/2002       | Kroczek         |       |          |                            |
|                       | AI        | 20020182667     | 12/05/2002       | Kroczek         |       |          |                            |

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| Examiner Initial  | Desig. ID | Document Number | Publication Date | Country or Patent Office | Class | Subclass | Translation |
|   |           |                 |                  |                          |       |          | Yes No      |
|   | AJ        | WO 95/33770     | 12/14/1995       | WIPO                     |       |          |             |
|   | AK        | WO 97/26912     | 07/31/1997       | WIPO                     |       |          |             |
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|   | AO        | JP 5-72204      | 03/23/1993       | Japan                    |       |          | Abstract    |
|   | AP        | JP 11-228442    | 08/24/1999       | Japan                    |       |          | Abstract    |
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|   | AR        | Abbas, "T-cell stimulation: an abundance of B7s," NAT MED. 5(12):1345-6 (1999)  |
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| J0   | AV        | Dong et al., "B7-H1, a third member of the B7 family, co-stimulates T-cell proliferation and interleukin-10 secretion," NAT. MED. 5(12):1365-9 (1999)   |
|  | AW        | Eljaschewitsch et al., "Identification of a novel activation antigen on human CD4+ T cells," IMMUNOBIOL., 194(1-3):27 (1995)  |
|  | AX        | Goding, "Monoclonal Antibodies: Principles and Practice," 2 <sup>nd</sup> Edition, Academic Press, Orlando, Florida, Chapter 8, pages 281-293 (1986)  |
|  | AY        | Goni et al., "Structural and idiotypic characterization of the L chains of human IgM autoantibodies with different specificities," J. Immunol. 142(9):3158-63 (1989)  |
|  | AZ        | Gonzalo et al., "ICOS is critical for T helper cell-mediated lung mucosal inflammatory responses," NAT IMMUNOL. 2(7):597-604 (2001)   |
|  | AAA       | Harlow and Lane, "Antibodies: A Laboratory Manual," Cold Spring Harbor Laboratory, page 285 (1988)  |
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|  | AGG       | Ling et al., "Differential expression of inducible costimulator-ligand splice variants: lymphoid regulation of mouse GL50-B and human GL50 molecules," J IMMUNOL. 166(12):7300-8 (2001)   |
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| Examiner Signature<br><i>Takashi Tsuji</i>   | Date Considered<br>11/07/2005 |
| EXAMINER: Initials citation considered. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |                               |

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| Substitute Form PTO-1449<br>(Modified)   | U.S. Department of Commerce<br>Patent and Trademark Office | Attorney's Docket No.<br><b>14539-006002</b> | Application No.<br><b>10/625,105</b> |
| <b>Information Disclosure Statement<br/>by Applicant</b><br>(Use several sheets if necessary)<br><br>(37 CFR §1.98(b)) |  | Applicant<br><b>Takashi Tsuji et al.</b>     |                                      |
|  |  | Filing Date<br><b>July 22, 2003</b>          | Group Art Unit                       |

| <b>Other Documents (include Author, Title, Date, and Place of Publication)</b> |           |   |
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| Examiner Initial   | Desig. ID | Document  |
| JO   | ARR       | Rottman et al., "The costimulatory molecule ICOS plays an important role in the immunopathogenesis of EAE," NAT IMMUNOL. 2(7):605-11 (2001)   |
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| EXAMINER: Initials citation considered. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |                                      |

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| Substitute Form PTO-1449<br><i>(Modified)</i><br>MAR 16 2005<br>U.S. Patent & Trademark Office                    | U.S. Department of Commerce<br>Patent and Trademark Office | Attorney's Docket No.<br>14539-006002 | Application No.<br>10/625,105 |
| <b>Information Disclosure Statement<br/>by Applicant</b><br>(Use several sheets if necessary)<br>(37 CFR 1.98(b)) |  | Applicant<br>Takashi Tsuji et al.     |                               |
|   |  | Filing Date<br>July 22, 2003          | Group Art Unit<br>1644        |

| U.S. Patent Documents |           |                 |                  |                   |       |          |                            |
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| Examiner Initial      | Desig. ID | Document Number | Publication Date | Patentee          | Class | Subclass | Filing Date If Appropriate |
| JO                    | AA        | 6,451,305       | 09/17/2002       | Boussiotis et al. | —     | —        | —                          |

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| Examiner Initial  | Desig. ID | Document Number | Publication Date | Country or Patent Office | Class | Subclass | Translation<br>Yes      No |
| JO  | AB        | 2,293,666       | 12/17/1998       | Canada                   | —     | —        | —                          |

| Other Documents (include Author, Title, Date, and Place of Publication) |           |          |  |  |  |  |  |
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| Examiner Initial  | Desig. ID | Document |  |  |  |  |  |
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| Examiner Signature<br><i>Elion Ansperger</i>  | Date Considered<br><i>11/07/2005</i> |
| EXAMINER: Initials citation considered. Drawline through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. |                                      |

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| Substitute Form PTO-1449<br><i>(Notated)</i><br>MAY 11 2005<br><i>Information Disclosure Statement</i><br>by Applicant<br>(Use several sheets if necessary)<br>(37 CFR 1.98(b)) | U.S. Department of Commerce<br>Patent and Trademark Office | Attorney's Docket No.<br><b>14539-006002</b> | Application No.<br><b>10/625,105</b> |
|   |  | Applicant<br><b>Takashi Tsuji et al.</b>     |                                      |
|   |  | Filing Date<br><b>July 22, 2003</b>          | Group Art Unit<br><b>1644</b>        |

| <b>U.S. Patent Documents</b> |           |                 |                  |          |       |          |                            |
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| Examiner Initial             | Desig. ID | Document Number | Publication Date | Patentee | Class | Subclass | Filing Date If Appropriate |
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| <b>Foreign Patent Documents or Published Foreign Patent Applications</b> |           |  |                  |                          |       |          |                            |
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| Examiner Initial   | Desig. ID | Document Number  | Publication Date | Country or Patent Office | Class | Subclass | Translation<br>Yes      No |
| <i>JO</i>  | AA        | WO 89/06138<br>(English language equivalent of RU 2047177) | 07/13/1989       | WIPO                     |       |          |                            |
| <i>JO</i>  | AB        | WO 94/11499  | 05/26/1994       | WIPO                     |       |          |                            |

| <b>Other Documents (include Author, Title, Date, and Place of Publication)</b> |           |          |
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| Examiner Signature<br><i>Dian Ouspensky</i>  | Date Considered<br><i>11/07/2005</i> |
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